

Amendment to the Claims

1-16. Canceled

17. (Previously presented) A method of delivering an active agent into the central nervous system of an animal comprising administering to said animal a conjugate comprising said agent conjugated to a Receptor Associated Protein (RAP) polypeptide consisting of an amino acid sequence at least 80% identical to amino acids 221-323 of RAP (SEQ ID NO: 1), wherein said RAP polypeptide retains megalin-binding activity and wherein said agent is delivered into the central nervous system.

18. (Previously presented) A method of increasing transcytosis of an active agent across the blood-brain barrier of an animal, comprising administering to said animal a conjugate comprising said agent conjugated to a Receptor Associated Protein (RAP) polypeptide consisting of an amino acid sequence at least 80% identical to amino acids 221-323 of RAP (SEQ ID NO: 1), wherein said RAP polypeptide retains megalin-binding activity and wherein said agent is transcytosed across the blood-brain barrier.

19. (Previously presented) A method of treating a disorder of the CNS in a mammal comprising administering to said mammal a conjugate comprising an effective amount of a therapeutic agent conjugated to a Receptor Associated Protein (RAP) polypeptide consisting of an amino acid sequence at least 80% identical to amino acids 221-323 of RAP (SEQ ID NO: 1).

20. Canceled

21. (Previously presented) The method of claim 19, wherein said disorder is selected from the group consisting of Huntington's Disease, Alzheimer's Disease, Parkinson's Disease, Multiple Sclerosis, Amyotrophic Lateral Sclerosis, ischemia-related disease and stroke, spinal muscular atrophy, cerebellar degeneration, perivenous encephalitis, schizophrenia, epilepsy and a central nervous system cancer.

22. (Withdrawn) The method of claim 21, wherein said disorder is a central nervous system cancer and said agent is a cancer chemotherapeutic agent.

23-57. Canceled

58. (Previously presented) The method of claim 17 or 18 wherein the animal is a human.

59. (Previously presented) The method of claim 58 wherein the human is suffering from a disorder selected from the group consisting of Huntington's Disease, Alzheimer's Disease, Parkinson's Disease, Multiple Sclerosis, Amyotrophic Lateral Sclerosis, ischemia-related disease and stroke, spinal muscular atrophy, cerebellar degeneration, perivenous encephalitis, schizophrenia, epilepsy and a central nervous system cancer.

60. (Previously presented) The method of claim 17 or 18 wherein the agent is a neurotrophic factor.

61. (Previously presented) The method of claim 17 or 18 wherein the therapeutic agent is a neurotrophic factor selected from the group consisting of Glial-Derived Neurotrophic Factor, Nerve Growth Factor, Brain-Derived Neurotrophic Factor, Neurotrophin-3, Neurotrophin-4/5, aFGF, bFGF, CNTF, Leukaemia Inhibitory Factor, Cardiotrophin-1, TGF β , BMPs, GDFs, Neurturin, Artemin, Persephin, EGF, TGF α , Neuregulins, IGF-1, IGF-2, ADNF and PDGF.

62. (Previously presented) The method of claim 17 or 18 wherein the therapeutic agent is brain-derived neurotrophic factor (BDNF).

63. (New) The method of any one of claims 17 to 19 wherein the polypeptide is at least 85% identical to amino acids 221-323 of RAP (SEQ ID NO: 1).

64. (New) The method of any one of claims 17 to 19 wherein the polypeptide is at least 90% identical to amino acids 221-323 of RAP (SEQ ID NO: 1).